

SEQUENCE LISTING

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SEP 2 6 2001 TECH CENTER 1600/2900

<110> Wang, Yi Mueller, John Matis, Louis A.

<120> Chimeric Proteins for Diagnosis and Treatment of Diabetes

<130> 109488-135

<140> US 09/528,225 <141> 2000-03-21

<150> PCT/US98/27408 <151> 1998-12-23

<150> US 60/068,648 <151> 1997-12-23

<160> 37

<170> PatentIn version 3.1

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<213> Artificial Sequence

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Arg Glu Ala Glu Asp Leu Asn Met Tyr Ala Met Met Ile Ala Arg Phe 35 40 45

Lys Met Phe Pro Glu Val Lys Glu Lys Gly Met Ala Ala Leu Pro Arg 50 55 60

Leu Ile Ala Phe Thr Ser Glu Lys Cys Leu Glu Leu Ala Glu Tyr Leu 65 70 75 80

Tyr Asn Ile Ile Lys Asn Arg Glu Gly Tyr Glu Met Val Phe Asp Gly 85 90 95

Lys Pro Gln His Thr Asn Val Cys Phe Trp Tyr Ile Pro Pro Ser Leu 100 105 110

Arg Thr Leu Glu Asp Asn Glu Glu Arg Met Ser Arg Leu Ser Lys Val

Ala Pro Val Ile Lys Ala Arg Met Met Glu Tyr Gly Thr Thr Met Val 130 135 140

Ser Tyr Gln Pro Leu Gly Asp Lys Val Asn His His His His His 145 150 155 160

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Arg Glu Ala Glu Asp Leu Met Asn Ile Leu Leu Gln Tyr Val Val Lys
35 40 45

Ser Phe Asp Asn Met Tyr Ala Met Met Ile Ala Arg Phe Lys Met Phe 50 55 60

Pro Glu Val Lys Glu Lys Gly Met Ala Ala Leu Pro Arg Leu Ile Ala 65 70 75 80

Phe Thr Ser Glu His Ser His Phe Ser Leu Lys Lys Cys Leu Glu Leu 85 90 95

Ala Glu Tyr Leu Tyr Asn Ile Ile Lys Asn Arg Glu Gly Tyr Glu Met
100 105 110

Val Phe Asp Gly Lys Pro Gln His Thr Asn Val Cys Phe Trp Tyr Ile 115 120 125

Pro Pro Ser Leu Arg Thr Leu Glu Asp Asn Glu Glu Arg Met Ser Arg 130 135 140

Leu Ser Lys Val Ala Pro Val Ile Lys Ala Arg Met Met Glu Tyr Gly
145 150 155 160

Thr Thr Met Val Ser Tyr Gln Pro Leu Gly Asp Lys Val Asn His His 165 170 175

His His His His 180

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Trp Leu Val Cys Gly Glu Arg Gly Phe Phe Tyr Thr Pro Lys Thr Arg 20 25 30

Arg Glu Ala Glu Asp Leu Met Asn Ile Leu Leu Gln Tyr Val Val Lys 35 40 45

Ser Phe Asp Asn Met Tyr Ala Met Met Ile Ala Arg Phe Lys Met Phe 50 55 60

Pro Glu Val Lys Glu Lys Gly Met Ala Ala Leu Pro Arg Leu Ile Ala 65 70 75 80

Phe Thr Ser Glu His Ser His Phe Ser Leu Lys Lys Cys Leu Glu Leu 85 90 95

Ala Glu Tyr Leu Tyr Asn Ile Ile Lys Asn Arg Glu Gly Tyr Glu Met 100 105 110

Val Phe Asp Gly Lys Pro Gln His Thr Asn Val Cys Phe Trp Tyr Ile 115 120 125

Pro Pro Ser Leu Arg Thr Leu Glu Asp Asn His His His His His 130 135 140

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Arg Glu Ala Glu Asp Leu Gln Val Gly Gln Val Glu Leu Gly Gly 35 40 45

Pro Gly Ala Gly Ser Leu Gl
n Pro Leu Ala Leu Glu Gly Ser Leu Gl
n 50 $$ 55 $$ 60

Lys Arg Gly Thr Asn Met Phe Thr Tyr Glu Ile Ala Pro Val Phe Val 65 70 75 80

Leu Leu Glu Tyr Val Thr Leu Lys Lys Met Arg Glu Ile Ile Gly Trp 85 90 95

Pro Gly Gly Ser Gly Asp Gly Gly Met Asn Ile Leu Leu Gln Tyr 100 105 110

Val Val Lys Ser Phe Asp Asn Met Tyr Ala Met Met Ile Ala Arg Phe 115 120 125

Lys Met Phe Pro Glu Val Lys Glu Lys Gly Met Ala Ala Leu Pro Arg 130 135 140

Leu Gly Gly Gly Ile Ala Phe Thr Ser Glu His Ser His Phe Ser Leu 145 150 155 160

Lys Lys Gly Ala Ala Leu Gly Ile Gly Thr Asp Ser Val Ile His 165 170 175

His His His His His 180

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Arg Glu Ala Glu Asp Leu Gln Val Gly Gln Val Glu Leu Gly Gly Gly 35 40 45

Pro Gly Ala Gly Ser Leu Gln Pro Leu Ala Leu Glu Gly Ser Leu Gln 50 55 60

Lys Arg Gly Thr Asn Met Phe Thr Tyr Glu Ile Ala Pro Val Phe Val 65 70 75 80

Leu Leu Glu Tyr Val Thr Leu Lys Lys Met Arg Glu Ile Ile Gly Trp \$85\$ 90 95

Pro Gly Gly Ser Gly Asp Gly Gly Met Asn Ile Leu Leu Gln Tyr 100 105 110

Val Val Lys Ser Phe Asp Asn Met Tyr Ala Met Met Ile Ala Arg Phe 115 120 125

Lys Met Phe Pro Glu Val Lys Glu Lys Gly Met Ala Ala Leu Pro Arg 130 135 140

Leu Gly Gly Gly Ile Ala Phe Thr Ser Glu His Ser His Phe Ser Leu 145 150 155 160

Lys Lys Gly Ala Ala Ala Leu Gly Ile Gly Thr Asp Ser Val Ile Gly
165 170 175

Gly Gly Tyr Ile Pro Pro Ser Leu Arg Thr Leu Glu Asp Asn Glu Glu
180 185 190

Arg Met Ser Arg Leu Ser Lys Val Ala Pro Val Ile Lys Ala Arg Met 195 200 205

Met Glu Tyr Gly Thr Thr Met Val Ser Tyr Gln Pro Leu Gly Asp Lys 210 215 220

Val Asn His His His His His 225 230

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- Arg Glu Ala Glu Asp Leu Gln Val Gly Gln Val Glu Leu Gly Gly 35 40 45
- Pro Gly Ala Gly Ser Leu Gln Pro Leu Ala Leu Glu Gly Ser Leu Gln 50 55 60
- Lys Arg Gly Thr Asn Met Phe Thr Tyr Glu Ile Ala Pro Val Phe Val 65 70 75 80
- Leu Leu Glu Tyr Val Thr Leu Lys Lys Met Arg Glu Ile Ile Gly Trp
 85 90 95
- Pro Gly Gly Ser Gly Asp Gly Gly Met Asn Ile Leu Leu Gln Tyr $100 \hspace{1.5cm} 105 \hspace{1.5cm} 110 \hspace{1.5cm}$
- Val Val Lys Ser Phe Asp Asn Met Tyr Ala Met Met Ile Ala Arg Phe 115 120 125
- Lys Met Phe Pro Glu Val Lys Glu Lys Gly Met Ala Ala Leu Pro Arg 130 $\,$ 135 $\,$ 140 $\,$
- Leu Gly Gly Gly Ile Ala Phe Thr Ser Glu His Ser His Phe Ser Leu 145 150 155 160
- Lys Lys Gly Ala Ala Ala Leu Gly Ile Gly Thr Asp Ser Val Ile Gly 165 170 175
- Gly Gly Ile Glu His Asp Pro Arg Met Pro Ala Tyr Ile Ala Thr Gln 180 185 190
- Gly Pro Leu Ser His Thr Ile Ala Asp Phe Trp Gln Met Val Trp Glu 195 200 205
- Ser Gly Cys Thr Val Ile Val Met Leu Thr Pro Leu Val Glu Asp Gly 210 215 220
- Val Lys Gln Cys Asp Arg Tyr Trp Pro Asp Glu Gly Ala Ser Leu Tyr 225 230 235 240
- His Val Tyr Glu Val Asn Leu Val Ser Glu His Ile Trp Cys Glu Asp \$245\$ \$250\$ \$255\$
- Phe Leu Val Arg Ser Phe Tyr Leu Lys Asn Val Gln Thr Gln Glu Thr

260 265 270

Arg Thr Leu Thr Gln Phe His Phe Leu Ser Trp Pro Ala Glu Gly Thr 275 280 285

Pro Ala Ser Thr Arg Pro Leu Leu Asp Phe Arg Arg Lys Val Asn Lys 290 295 300

Cys Tyr Arg Gly Arg Ser Cys Pro Ile Ile Val His Cys Ser Asp Gly 305 310 315 320

Ala Gly Arg Thr Gly Thr Tyr Ile Leu Ile Asp Met Val Leu Asn Arg 325 330 335

Met Ala Lys Gly Val Lys Glu Ile Asp Ile Ala Ala Thr Leu Glu His 340 345 350

Val Arg Asp Gln Arg Pro Gly Leu Val Arg Ser Lys Asp Gln Phe Glu 355 360 365

Phe Ala Leu Thr Ala Val Ala Glu Glu Val Asn Ala Ile Leu Lys Ala 370 375 380

Leu Pro Gln His His His His His 385 390

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Tyr Leu Val Cys Gly Glu Arg Gly Phe Phe Tyr Thr Pro Lys Thr Arg 20 25 30

Arg Glu Ala Glu Asp Leu Gln Val Gly Gln Val Glu Leu Gly Gly

35 40 45

Pro Gly Ala Gly Ser Leu Gln Pro Leu Ala Leu Glu Gly Ser Leu Gln 50 55 60

Lys Arg Gly Thr Asn Met Phe Thr Tyr Glu Ile Ala Pro Val Phe Val 65 70 75 80

Leu Leu Glu Tyr Val Thr Leu Lys Lys Met Arg Glu Ile Ile Gly Trp 85 90 95

Pro Gly Gly Ser Gly Asp Gly Gly Gly Met Asn Ile Leu Leu Gln Tyr 100 105 110

Lys Met Phe Pro Glu Val Lys Glu Lys Gly Met Ala Ala Leu Pro Arg 130 135 140

Leu Gly Gly Gly Ile Ala Phe Thr Ser Glu His Ser His Phe Ser Leu 145 150 155 160

Lys Lys Gly Ala Ala Ala Leu Gly Ile Gly Thr Asp Ser Val Ile Gly 165 170 175

Gly Gly Tyr Ile Pro Pro Ser Leu Arg Thr Leu Glu Asp Asn Glu Glu 180 185 190

Arg Met Ser Arg Leu Ser Lys Val Ala Pro Val Ile Lys Ala Arg Met 195 200 205

Met Glu Tyr Gly Thr Thr Met Val Ser Tyr Gln Pro Leu Gly Asp Lys 210 215 220

Val Asn Gly Gly Gly Ile Glu His Asp Pro Arg Met Pro Ala Tyr Ile 225 230 235 240

Ala Thr Gln Gly Pro Leu Ser His Thr Ile Ala Asp Phe Trp Gln Met 245 250 255

Val Trp Glu Ser Gly Cys Thr Val Ile Val Met Leu Thr Pro Leu Val 260 265 270

Glu Asp Gly Val Lys Gln Cys Asp Arg Tyr Trp Pro Asp Glu Gly Ala 280 275 Ser Leu Tyr His Val Tyr Glu Val Asn Leu Val Ser Glu His Ile Trp 295 Cys Glu Asp Phe Leu Val Arg Ser Phe Tyr Leu Lys Asn Val Gln Thr 310 Gln Glu Thr Arg Thr Leu Thr Gln Phe His Phe Leu Ser Trp Pro Ala 330 325 Glu Gly Thr Pro Ala Ser Thr Arg Pro Leu Leu Asp Phe Arg Arg Lys 345 340 Val Asn Lys Cys Tyr Arg Gly Arg Ser Cys Pro Ile Ile Val His Cys 365 360 355 Ser Asp Gly Ala Gly Arg Thr Gly Thr Tyr Ile Leu Ile Asp Met Val 370 375 Leu Asn Arg Met Ala Lys Gly Val Lys Glu Ile Asp Ile Ala Ala Thr 395 400 390 Leu Glu His Val Arg Asp Gln Arg Pro Gly Leu Val Arg Ser Lys Asp 410 405 Gln Phe Glu Phe Ala Leu Thr Ala Val Ala Glu Glu Val Asn Ala Ile 425 Leu Lys Ala Leu Pro Gln His His His His His 440 435

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Arg Glu Ala Glu Asp Leu Gln Val Gly Gln Val Glu Leu Gly Gly Gly 35 40 45

Pro Gly Ala Gly Ser Leu Gln Pro Leu Ala Leu Glu Gly Ser Leu Gln 50 55 60

Lys Arg Gly Met Asn Ile Leu Leu Gln Tyr Val Val Lys Ser Phe Asp 65 70 75 80

Asn Met Tyr Ala Met Met Ile Ala Arg Phe Lys Met Phe Pro Glu Val 85 90 95

Lys Glu Lys Gly Met Ala Ala Leu Pro Arg Leu Ile Ala Phe Thr Ser 100 105 110

Glu His Ser His Phe Ser Leu Lys Lys Cys Leu Glu Leu Ala Glu Tyr 115 120 125

Leu Tyr Asn Ile Ile Lys Asn Arg Glu Gly Tyr Glu Met Val Phe Asp 130 135 140

Gly Lys Pro Gln His Thr Asn Val Cys Phe Trp Tyr Ile Pro Pro Ser 145 150 155 160

Leu Arg Thr Leu Glu Asp Asn His His His His His His 165 170

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cataco	caacg tttgcttttg gtacatcccg ccgagcctgc gtaccctgga agataacgaa	360
gaacgo	catga gccgtctgtc taaagttgcc ccggttatta aagcgcgcat gatggaatat	420
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Artificial Sequence

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